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<u>#1</u>	Search CB/ior-CEA.1	17:33:28	<u>0</u>

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                 patents
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L2 ANSWER 1 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN AN 1996:85276 CAPLUS

DN 124:172974

- TI Bacterial expression and characterization of a modified scFv fragment from the anti-carcinoembryonic antigen monoclonal antibody CB.

  CEA.1
- AU Vazquez, Javier E.; Perez, Lilia; Ayala, Martha; Canaan-Haden, Leonardo; De Lalla, C.; Sidoli, A.; Gavilondo, Jorge
- CS Center Genetic Engineering and Biotechnology, Havana, 10600, Cuba
- SO Biotecnologia Aplicada (1995), 12(2), 96-7 CODEN: BTAPEP; ISSN: 0864-4551
- PB Sociedad Iberolatinoamericana de Biotecnologia Aplicada a la Salud
- DT Journal
- LA English
- AB The authors report here the cloning, bacterial expression (Escherichia coli), and mol. characterization of an anti-carcinoembryonic antigen (CEA) single-chain Fv antibody fragment with potential use in the localization of tumors in CEA-pos. carcinoma patients.
- L2 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN
- AN 1993:78955 CAPLUS
- DN 118:78955
- TI Bacterial single-chain antibody fragments, specific for carcinoembryonic antigen
- AU Ayala, M.; Duenas, M.; Santos, A.; Vazquez, J.; Menendez, A.; Silva, A.; Gavilondo, J. V.
- CS Cent. Genet. Eng. Biotechnol., Cuba
- SO BioTechniques (1992), 13(5), 790-2, 794-6, 798-9 CODEN: BTNQDO; ISSN: 0736-6205
- DT Journal
- LA English
- AB Single-chain antibody (scFv) fragments were produced in bacteria specific for carcinoembryonic antigen (CEA). Polymerase chain reaction (PCR) was used for the cloning and modification of the heavy and light variable regions (VH and VL) of the mouse monoclonal antibody (MAb) CB-CEA.1. Culture supernatant, bacteria pellet and periplasm prepns. were assayed in Western blot, and a protein of .apprx.27 kDa was identified with rabbit antibodies specific for the Fab of CB-CEA.1. Bacterial supernatant and periplasm prepns. also inhibited the recognition of CEA by HRP-labeled CB-CEA.1 in ELISA. Periplasm prepns. were purified by affinity chromatog. with specific anti-idiotypic monoclonal antibodies. The Western blot of the eluates identified a protein of .apprx.27 kDa that blocked the recognition of CEA by HRP-labeled CB-CEA.
- L2 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN
- AN 1992:122152 CAPLUS

1 in ELISA.

- DN 116:122152
- TI Primer design for the cloning of immunoglobulin heavy-chain leader-variable regions from mouse hybridoma cells using the PCR
- AU Coloma, Maria J.; Larrick, James W.
- CS Genelabs, Inc., Redwood City, CA, 94063, USA
- SO BioTechniques (1991), 11(2), 152-4, 156 CODEN: BTNQDO; ISSN: 0736-6205
- DT Journal
- LA English
- AB To facilitate the rapid cloning and sequencing of rearranged murine heavy-chain variable regions, a set of universal primers was designed using conserved sequences of leader (signal peptide), framework one and constant regions of the Ig heavy-chain genes. RNA was extracted from the mouse hybridoma cells secreting monoclonal antibodies: IOR-T3 (anti-CD3), C6 (anti-Pl of N. meningitidis B385), IOR-T1 (anti-CD6), CB-CEA.1 (anti-carcinoembryonic antigen), CB-Fib.1 (anti-human fibrin) and CB-Hep.2 (anti-hepatitis B surface antigen). First-strand cDNA was synthesized and amplified using PCR. The primers successfully amplified correct size fragments from cDNA prepared from all

hybridomas. These methods will facilitate the cloning and sequencing of mouse Ig variable regions.

- L2 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN
- AN 1991:56891 CAPLUS
- DN 114:56891
- TI Specific amplification of rearranged immunoglobulin variable region genes from mouse hybridoma cells
- AU Gavilondo-Cowley, Jorge V.; Coloma, Maria J.; Vazquez, Javier; Ayala, Marta; Macias, Amparo; Fry, Kirk E.; Larrick, James W.
- CS Div. Hybridomas Anim. Models, Cent. Genet. Eng. Biotechnol., Havana, Cuba
- SO Hybridoma (1990), 9(5), 407-17 CODEN: HYBRDY; ISSN: 0272-457X
- DT Journal
- LA English
- This article describes how the polymerase chain reaction (PCR) and primers AB designed for conserved sequences of leader (L), framework one (FR1) and constant (CONST) regions of Ig light and heavy chain genes can be used for the cloning and sequencing of rearranged antibody variable regions from mouse hybridoma cells. RNA was extracted from the mouse hybridoma cells secreting MAbs: IOR-T3a (anti-CD3), C6 (anti-P1 of Neisseria meningitidis B385), IOR-T1 (anti-CD6), CB-CEA.1 (anti-carcinoembryonic antigen), and CB-Fib.1 (anti-human fibrin). First strand cDNA was synthesized and amplified using PCR. The newly designed primers are superior to others reported recently in the literature. Isolated PCR DNA fragments of C6 and IOR-T3a were sequenced after asym. amplification, or M13 cloning. The FR1/CONST primer combinations selectively amplified mouse light chains of groups kappa II, V, and VI, and heavy chains of groups IIa and IIc. The L/CONST primers for light chains amplified light chains from all 4 hybridomas. The methods greatly facilitate structural and functional studies of antibodies by reducing the efforts required to clone and sequence their variable regions.
- L2 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN
- AN 1990:156288 CAPLUS
- DN 112:156288
- TI Relative positions of some epitopes on carcinoembryonic antigen
- AU Larsson, Aake; Ghosh, Rahul; Hammarstroem, Sten
- CS Dep. Immunol., Univ. Stockholm, Stockholm, S-10691, Swed.
- SO Cancer Immunology Immunotherapy (1989), 30(2), 92-6 CODEN: CIIMDN; ISSN: 0340-7004
- DT Journal
- LA English
- AB To investigate whether anti-(carcinoembryonic antigen) monoclonal antibodies (mAb) react with single or repeated epitopes, sandwich RIAs in homologous and heterologous combinations were performed. Four mAb (I-27, I-47, II-17, and to some degree II-16) gave homologous binding, while two mAb (I-38S1 and II-10) did not. All these mAb except II-10 react with repeated epitopes. The relative positions of the epitopes recognized by these mAb and of three addnl. mAb (II-6, II-7, and CB-CEA-1) were investigated using a plate antibody competition test with enzyme-labeled carcinoembryonic antigen (CEA). The mAb I-38S1, II-6, II-7, II-10, II-16, and CB-CEA-

1 were mutually cross-reactive, and were classified as belonging to one epitope group. The mAb I-27 and I-47 fell outside this group and did not interfere with the binding of CEA conjugate to mAb II-17 either. They, therefore, represent a second epitope group. The mAb II-17 showed no interference with the binding of CEA to any of the other mAb and must therefore represent a third epitope group. The slopes of the plate antibody competition curves were used for calcn. of a correlation matrix, which in turn was used to depict the relative positions of the epitopes recognized by the mAb in the large group.

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09428616 PMID: 1418981

Bacterial single-chain antibody fragments, specific for carcinoembryonic antigen.

Ayala M; Duenas M; Santos A; Vazquez J; Menendez A; Silva A; Gavilondo J $_{J}$ 

Division of Immunotechnology and Diagnostics, Center for Genetic Engineering and Biotechnology, La Habana, Cuba.

BioTechniques (UNITED STATES) Nov 1992, 13 (5) p790-9, ISSN 0736-6205--Print Journal Code: 8306785

Publishing Model Print

Document type: Journal Article

Languages: ENGLISH

Main Citation Owner: NLM

Record type: MEDLINE; Completed

Subfile: INDEX MEDICUS

We have produced single-chain antibody (scFv) fragments in bacteria specific for carcinoembryonic antigen (CEA). Polymerase chain reaction (PCR) was used for the cloning and modification of the heavy and light variable regions (VH and VL) of the mouse monoclonal antibody (MAb) CB-CEA.1. A 14-amino acid linker was used in the synthesis of the scFv gene. The VH and VL regions were amplified from cDNA by PCR using 5' end FR1 and 3' end constant region primers, and then sequenced. VH was then amplified by PCR using an exact 5' end FR1 primer, and a phosphorylated (PP) 3' end primer for J2 that also encoded the first 7 amino acids of the linker. VL was amplified with a PP 5' end primer for FR1, also encoding the remaining 7 amino acids of the linker, and a 3' end primer for J5, plus a stop codon and a BqlII restriction site. The fragments were ligated and reamplified with the PP VH 5' and VL 3' end primers. The VH-linker-VL structure was blunt-cloned into expression vectors bearing the tryptophan promoter and pelB or ompA signal peptide sequences. Culture supernatant, bacteria pellet and periplasm preparations were assayed in Western blot and a protein of about 27 kDa was identified with rabbit antibodies specific for the Fab of CB-CEA.1. Bacterial supernatant and periplasm preparations inhibited the recognition of CEA by HRP-labeled CB-CEA.1 in enzyme-linked immunosorbent assay (ELISA). Periplasm preparations were purified by affinity chromatography with specific anti-idiotypic MAbs. The Western blot of the eluates identified a protein of approximately 27 kDa that blocked the recognition of CEA by HRP-labeled CB-CEA.1 in ELISA. The VH-linker-VL structure was cloned into a vector bearing the lacZ promoter and the pelB signal peptide. The recombinant bacterial clones also expressed about 27 kDa scFv, specific for CEA.

Descriptors: \*Antibodies, Neoplasm--immunology--IM; \*Antibody Specificity --genetics--GE; \*Carcinoembryonic Antigen--immunology--IM; \*Immunoglobulin Fragments--immunology--IM; Amino Acid Sequence; Animals; Antibodies, Antibodies, Neoplasm--genetics--GE; Monoclonal -- immunology -- IM; Cloning, Chromatography, Affinity; Molecular; Sequence; Enzyme-Linked Immunosorbent Assay; Escherichia coli; Single-Stranded; Genetic Vectors; Hybridomas; Immunoglobulin Fragments--genetics--GE; Mice; Molecular Sequence Data; Plasmids; Polymerase Chain Reaction; Protein Binding; Recombinant Proteins--metabolism--ME

Molecular Sequence Databank No.: GENBANK/D10270; GENBANK/D10271; GENBANK/D10272; GENBANK/D10273; GENBANK/D10274; GENBANK/D10275; GENBANK/D10276; GENBANK/M63506; GENBANK/S49453; GENBANK/S49457

CAS Registry No.: 0 (Antibodies, Monoclonal); 0 (Antibodies, Neoplasm); 0 (Carcinoembryonic Antigen); 0 (DNA, Single-Stranded); 0 (Immunoglobulin Fragments); 0 (Recombinant Proteins)

Gene Symbol: scFv

Record Date Created: 19921210

Record Date Completed: 19921210

5/9/2 (Item 2 from file: 155)
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09010334 PMID: 1931008
Primer design for the cloning of

Primer design for the cloning of immunoglobulin heavy-chain leader-variable regions from mouse hybridoma cells using the PCR.

Coloma M J; Larrick J W; Ayala M; Gavilondo-Cowley J V

Genelabs, Inc., Redwood City, CA 94063.

BioTechniques (UNITED STATES) Aug 1991, 11 (2) p152-4, 156, ISSN 0736-6205--Print Journal Code: 8306785

Publishing Model Print

Document type: Journal Article; Research Support, Non-U.S. Gov't

Languages: ENGLISH

Main Citation Owner: NLM

Record type: MEDLINE; Completed

Subfile: INDEX MEDICUS

To facilitate the rapid cloning and sequencing of rearranged murine heavy-chain variable regions, we have designed a set of universal primers using conserved sequences of leader (signal peptide), framework one and constant regions of the immunoglobulin heavy-chain genes. RNA was extracted from the mouse hybridoma cells secreting monoclonal antibodies: IOR-T3 (anti-CD3), C6 (anti-P1 of N. meningitidis B385), IOR-T1 (anti-CD6), CB-CEA.1 (anti-carcinoembryonic antigen), CB-Fib.1 (anti-human fibrin) and CB-Hep.2 (anti-hepatitis B surface antigen). First-strand cDNA was synthesized and amplified using PCR. The primers successfully amplified correct size fragments from cDNA prepared from all hybridomas. These methods will facilitate the cloning and sequencing of mouse immunoglobulin variable regions.

Descriptors: \*Immunoglobulin Heavy Chains--genetics--GE; \*Immunoglobulin Variable Region--genetics--GE; \*Polymerase Chain Reaction; Animals; Base Sequence; Cell Line; Cloning, Molecular; DNA--chemical synthesis--CS; Hybridomas; Mice; Molecular Sequence Data; Polydeoxyribonucleotides --chemical synthesis--CS

CAS Registry No.: 0 (Immunoglobulin Heavy Chains); 0 (Immunoglobulin Variable Region); 0 (Polydeoxyribonucleotides); 9007-49-2 (DNA)

Record Date Created: 19911203
Record Date Completed: 19911203

5/9/3 (Item 3 from file: 155)
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08323758 PMID: 2611022

CEA in colonic adenocarcinomas and precancerous lesions. An immunohistochemical study with a novel monoclonal antibody.

Tormo B R; Gavilondo J V; Dominguez C; Freyre M; Rodriguez T; Biberfeld P Instituto Nacional de Oncologia y Radiobiologia, Departmento de Biologia, La Habana, Cuba.

APMIS - acta pathologica, microbiologica, et immunologica Scandinavica (DENMARK) Dec 1989, 97 (12) p1073-80, ISSN 0903-4641--Print Journal Code: 8803400

Publishing Model Print

Document type: Journal Article; Research Support, Non-U.S. Gov't

Languages: ENGLISH

Main Citation Owner: NLM

Record type: MEDLINE; Completed

Subfile: INDEX MEDICUS

Using a new anti-CEA (CB-CEA-1) murine monoclonal antibody, the expression of carcinoembryonic antigen (CEA) was studied in normal, premalignant and malignant human adult tissues with particular emphasis on colorectal mucosa. The CB-CEA-1 epitope was poorly expressed in normal adult tissues but was consistently found in colon cancers and adenomas in distinctive immunohistochemical patterns. Some apical staining was found with CB-CEA-1 in cells of normal colon mucosa whereas colon adenocarcinomas had a predominantly cytoplasmic staining pattern. Colonic adenomas presented a varied staining pattern. Some showed apical staining, others a CEA distribution pattern similar to that of adenocarcinomas, particularly those with a villous component. Our findings indicate a differential expression of CB-CEA-1 in adenoma cells in relation to their potential for malignant transformation. The possible usefulness of this Mab defined epitope for diagnostic and therapeutic purposes is indicated.

Descriptors: \*Adenocarcinoma--metabolism--ME; \*Carcinoembryonic Antigen --metabolism--ME; \*Colonic Neoplasms--metabolism--ME; \*Precancerous Conditions--metabolism--ME; Adenocarcinoma--pathology--PA; Antibodies, Monoclonal--diagnostic use--DU; Antibodies, Monoclonal--immunology--IM; Carcinoembryonic Antigen--immunology--IM; Colonic Neoplasms--diagnosis--DI; Colonic Neoplasms--pathology--PA; Humans; Immunohistochemistry; Precancerous Conditions--diagnosis--DI; Precancerous Conditions--pathology--PA

CAS Registry No.: 0 (Antibodies, Monoclonal); 0 (Carcinoembryonic Antigen)

Record Date Created: 19900302 Record Date Completed: 19900302

5/9/4 (Item 4 from file: 155)
DIALOG(R)File 155:MEDLINE(R)

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08300833 PMID: 2480845

Relative positions of some epitopes on carcinoembryonic antigen.

Larsson A; Ghosh R; Hammarstrom S

Department of Immunology, University of Stockholm, Sweden.

Cancer immunology, immunotherapy - CII (GERMANY, WEST) 1989, 30 (2) p92-6, ISSN 0340-7004--Print Journal Code: 8605732

Publishing Model Print

Document type: Journal Article; Research Support, Non-U.S. Gov't

Languages: ENGLISH

Main Citation Owner: NLM

Record type: MEDLINE; Completed

Subfile: INDEX MEDICUS

anti-(carcinoembryonic antigen) monoclonal investigate whether react with single or repeated epitopes, sandwich (mAb) antibodies homologous and heterologous combinations were radioimmunoassays in performed. Four mAb (I-27, I-47, II-17 and to some degree II-16) gave homologous binding while two mAb (I-38S1 and II-10) did not. Taken together with previous immunoprecipitation studies we conclude that all these mAb except II-10 react with repeated epitopes. The relative positions of the epitopes recognized by these mAb and of three additional mAb (II-6, II-7 and CB-CEA-1) were investigated using a plate antibody competition test with enzyme-labelled carcinoembryonic antigen (CEA). mAb I-38S1, II-6, II-7, II-10, II-16 and CB-CEA-1 were mutually cross-reactive, and were classified as belonging to one epitope group. mAb I-27 and I-47 fell outside this group and did not interfere with the binding of CEA conjugate to mAb II-17 either. They therefore represent a second epitope group. mAb

```
II-17 showed no interference with the binding of CEA to any of the other
mAb and must therefore represent a third epitope group. The slopes of the
                  competition curves were used for calculation of a
      antibody
correlation matrix, which in turn was used to depict the relative positions
of the epitopes recognized by the mAb in the large group.
 Descriptors:
                  *Carcinoembryonic
                                    Antigen--immunology--IM;
--analysis--AN; Antibodies, Monoclonal--diagnostic use--DU; Humans
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             (Epitopes)
Antigen); 0
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          BIOSIS NO.: 199698635086
 Bacterial expression and characterization of a modified scFv fragment from
 the anti-carcinoembryonic antigen monoclonal antibody CB.CEA.1
AUTHOR: Vazquez Javier E (Reprint); Perez Lilia (Reprint); Ayala Martha
  (Reprint); Canaan-Haden Leonardo (Reprint); De Lalla C; Sidoli A;
  Gavilondo Jorge (Reprint)
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  Cuba**Cuba
JOURNAL: Biotecnologia Aplicada 12 (2): p96-97 1995 1995
DOCUMENT TYPE: Article
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LANGUAGE: English
DESCRIPTORS:
  MAJOR CONCEPTS: Biochemistry and Molecular Biophysics; Infection;
    Metabolism; Oncology--Human Medicine, Medical Sciences; Physiology
  BIOSYSTEMATIC NAMES: Hominidae--Primates, Mammalia, Vertebrata, Chordata,
    Animalia
  ORGANISMS: human (Hominidae)
  COMMON TAXONOMIC TERMS: Animals; Chordates; Humans; Mammals; Primates;
    Vertebrates
                        ANTIBODY RECOGNITION PATTERN; CARCINOEMBRYONIC
  MISCELLANEOUS TERMS:
    ANTIGEN; MOLECULAR CHARACTERIZATION; NOTE; TUMOR LOCALIZATION
CONCEPT CODES:
  10064 Biochemistry studies - Proteins, peptides and amino acids
  10068 Biochemistry studies - Carbohydrates
  10506 Biophysics - Molecular properties and macromolecules
  13004 Metabolism - Carbohydrates
  13012 Metabolism - Proteins, peptides and amino acids
  24006 Neoplasms - Biochemistry
  31000 Physiology and biochemistry of bacteria
  36002 Medical and clinical microbiology - Bacteriology
BIOSYSTEMATIC CODES:
  86215 Hominidae
S (CB (W) CEA (W) 1)
          442858 CB
           56221 CEA
        16945423 1
              20 (CB (W) CEA (W) 1)
S S6 AND (THERAP? OR TREAT?)
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DIALOG(R) File 155: MEDLINE(R)
(c) format only 2007 Dialog. All rts. reserv.
08323758
          PMID: 2611022
 CEA
           colonic adenocarcinomas
                                       and
                                             precancerous
 immunohistochemical study with a novel monoclonal antibody.
 Tormo B R; Gavilondo J V; Dominguez C; Freyre M; Rodriguez T; Biberfeld P
  Instituto Nacional de Oncologia y Radiobiologia, Departmento de Biologia,
La Habana, Cuba.
  APMIS - acta pathologica, microbiologica, et immunologica Scandinavica (
          Dec 1989, 97 (12) p1073-80, ISSN 0903-4641--Print
DENMARK)
Journal Code: 8803400
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 Languages: ENGLISH
 Main Citation Owner: NLM
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             new anti-CEA (CB-CEA-1) murine monoclonal antibody,
  Using
expression of carcinoembryonic antigen (CEA) was studied in normal,
premalignant and malignant human adult tissues with particular emphasis on
colorectal mucosa. The CB-CEA-1 epitope was poorly expressed in normal
adult tissues but was consistently found in colon cancers and adenomas in
distinctive immunohistochemical patterns. Some apical staining was found
with CB-CEA-1 in cells of normal colon mucosa whereas colon adenocarcinomas
         predominantly cytoplasmic staining pattern. Colonic adenomas
presented a varied staining pattern. Some showed apical staining, others a
CEA distribution pattern similar to that of adenocarcinomas, particularly
those with a villous component. Our findings indicate a differential
expression of CB-CEA-1 in adenoma cells in relation to their potential for
malignant transformation. The possible usefulness of this Mab defined
epitope for diagnostic and therapeutic purposes is indicated.
  Descriptors: *Adenocarcinoma--metabolism--ME; *Carcinoembryonic Antigen
                                Neoplasms--metabolism--ME; *Precancerous
--metabolism--ME;
                    *Colonic
Conditions -- metabolism -- ME;
                              Adenocarcinoma -- pathology -- PA; Antibodies,
Monoclonal--diagnostic use--DU; Antibodies, Monoclonal--immunology--IM;
Carcinoembryonic Antigen--immunology--IM; Colonic Neoplasms--diagnosis--DI;
            Neoplasms--pathology--PA;
                                          Humans;
                                                     Immunohistochemistry;
Precancerous Conditions--diagnosis--DI; Precancerous Conditions--pathology
- - PA
                            (Antibodies, Monoclonal); 0 (Carcinoembryonic
  CAS Registry No.: 0
Antigen)
  Record Date Created: 19900302
  Record Date Completed: 19900302
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S6	20	(CB (W) CEA (W) 1)
<b>S</b> 7	4	S6 AND (THERAP? OR TREAT?)
S8	1	RD S7 (unique items)
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